

Supplementary file 1 – Matrix of evidence utilised in developing the provisional messages for Stage 2, phases 1 and 2 patient interviews

Draft messages Developed in HPV Core Messages Study (marked *) or added following expert group discussion to address non-cervical cancers	Qualitative interview & postal survey evidence For messages developed in HPV core messages study	Systematic review evidence from core messages study Underpinning evidence from a review of reviews ----- Evidence of information needs from reviews of views on HPV vaccination and testing	Evidence from other published sources Identified in the 'Talking about HPV' systematic review, or targeted searches, or by members of the expert group	Telephone interviews with healthcare professionals Summarised comments and illustrative quotes	Provisional messages after expert group review
HPV is very common; most sexually active people will have it at some point in their life.*	<p>Most women in the NHS cervical screening programme HPV pilot sites who were interviewed said they had been unaware of HPV prior to their results letter.</p> <p>Only 9.2 % of the public and 37.3% of healthcare professionals surveyed thought the statement, "Most people will get an HPV infection during their life" was true.</p> <p>6.1% of the public and 2.5% of healthcare professionals believed HPV infection to be rare.</p>	<p>The prevalence of HPV in the USA ranged from 14% to more than 90%.¹</p> <p>The estimated global HPV prevalence in women with normal cytological findings was 11.7%.²</p> <p>-----</p> <p>Few knew HPV is highly prevalent.³</p>	<p>Nearly 3 out of 4 people between the ages of 15 and 49 have been have been infected with HPV at some point in their lives.⁴</p> <p>Most of us will acquire a HPV infection at some point in our lives, usually without knowing about it. Telling patients this, and being inclusive, will help them feel that HPV is 'normal'.⁵</p> <p>Most sexually active people will get HPV early in their lifetimes.⁶</p>	<p>Clinicians explain to patients that HPV infection is very common and can take years to develop, but often seek to avoid emphasising the sexually transmitted nature of the virus.</p> <p>"It is as common as having a cold"</p> <p>"I explain that the virus is like the flu virus in terms of it is a virus and that pretty much all of us can get it...I hardly ever say that it is sexually transmitted unless they bring that up.</p> <p>"We try and say to them it is a virus that's sexually transmitted but not in a way to make them feel they have done something wrong..."</p> <p>"I tell them sex is a healthy thing and that most of us are carrying it (HPV) – so they don't feel dirty"</p> <p>"A lot of men carry it but it's not like having chlamydia"</p>	<p>1. HPV is a very common virus that most sexually active people will be exposed to at some point in their life.</p>
HPV is just as common in men as in women.*	<p>In interviews, parents of vaccination-aged girls were unsure of the value of the vaccination programme of boys were not to be vaccinated to help stop the spread of HPV.</p>	<p>HPV infection is highly prevalent in sexually active men and can be detected by a variety of specimens and methods.⁷</p> <p>-----</p> <p>The term "high-risk" was poorly understood; some women associated it with risky behaviour.⁸</p> <p>The review suggests that HPV is a risk factor for some head and neck cancers.⁹</p>	<p>Partners tend to share HPV...⁶</p>	<p>"A lot of men carry it but it's not like having chlamydia"</p>	<p>2. HPV is common in both men and women and is often shared between partners.</p> <p><i>Deselected after consultation with expert group</i></p>
Some types of HPV are 'high risk' and can cause abnormal cell changes that could lead, eventually, to cancer.*	<p>"High-risk" was not a term used by interview participants – the consideration is that protection is offered against the viruses that cause cervical cancer, not that there are other HPVs that are not linked with cervical cancer.</p>	<p>-----</p> <p>The term "high-risk" was poorly understood; some women associated it with risky behaviour.⁸</p> <p>The review suggests that HPV is a risk factor for some head and neck cancers.⁹</p>	<p>'Most cancers of the vagina and anus are likewise caused by HPV, as are a fraction of cancers of the vulva, penis, and oropharynx. HPV-16 and -18 account for about 70% of</p>	<p>Patients ask about the other risks of HPV exposure and that other cancers can come as a result of HPV.</p>	<p>3. Infection with certain HPV types causes all cervical cancer,</p>
Infection with HPV types that can cause cervical cancer can also cause other	<p>Women interviewed in the NHS cervical screening programme HPV pilot sites questioned whether cancers other than cervical cancer</p>				

<p>cancers such as <i>penile, vulvar, vaginal, anal and head and neck.*</i></p>	<p>can be caused by HPV, particularly whether men are more at risk from other HPV related cancers.</p>	<p>The overall HPV prevalence was 25.9% in head and neck cancers.¹⁰</p> <p>Oral infection with HPV is a significant independent risk factor for oral squamous cell carcinoma.¹¹</p> <p>The direction of effect observed was towards an increased risk of laryngeal squamous cell carcinoma when HPV was detected.¹²</p> <p>-----</p> <p>Some women questioned whether HPV can cause other cancers.⁸</p>	<p>cancers of the cervix, vagina, and anus and for about 30–40% of cancers of the vulva, penis, and oropharynx.¹³</p> <p>Estimated annual new HPV-related cancers globally: 10,500 penile; 27,300 anal; 19,960 vulvar and vaginal; 529,800 cervical; 60,000 head and neck¹³⁻¹⁷</p>	<p>“It’s bad enough saying you contracted the virus via your cervix, but via your mouth is a bit much sometimes.”</p> <p>“It may come up (in the case of females) that they have had cervical screening and that the virus that has caused the change in the cervix has done the same in the anus.”</p> <p>“For cervical cancer it’s acknowledged about sexual activity, that it is the HPV virus but anal, vulvar and vaginal cancers have not had that opportunity. Head & Neck has become more relevant now that it’s been associated with HPV infection and also with better prognosis, so you’re giving people potentially useful information whereas in anal cancer is it useful to give information around this as standard or are we just raising another issue which doesn’t really help them?”</p> <p>Clinicians explain to patients that most people’s immune systems will get rid of HPV within a year or two.</p> <p>“It is quite difficult to explain to people that you can go decades between getting infected with the virus and these cancers happening is quite a lot for people to get their head round.”</p> <p>“The important message to get over to patients is that we are all exposed to viruses during normal sexual or social interaction and that their immune system has not knocked this one out...and a whole series of things have happened since which have led to you having a tumour.”</p> <p>Clinicians explain that HPV can stay in the body dormant and not cause problems for years and years.</p> <p>“I told a patient that he may have picked up this infection 10-20 years ago and that everyone is exposed to it. He started crying and said he was glad I had told him about it because his wife was blaming herself that she gave him this cancer....”</p>	<p>almost all anal cancer and can also cause some other cancers such as penile, vulvar, vaginal, and head and neck</p>
<p>HPV can usually be cleared by the immune system but occasionally, it stays in the cells of the cervix for several years. This persistent infection can cause the abnormalities that could lead to cervical cancer.*</p>	<p>There was poor understanding of how an HPV infection can lead to a cervical cancer. Simple and clear information was wanted, by Healthcare professionals and women who were HPV-positive, about the process of development of cervical cancer.</p> <p>Only 4% of members of the public surveyed believed the statement “HPV infection usually goes away without needing treatment” to be true, compared with 35% of healthcare professionals.</p>	<p>HPV persistence was consistently and strongly associated with CIN2-3.¹⁸</p>	<p>A healthy immune system suppresses the virus in most infected persons.⁶</p> <p>The vast majority of HPV infections are transient and most healthy individuals will effectively clear the virus. However, there may be significant person-to-person variability in the time required to complete viral clearance. In most people, HPV is cleared within a year or two.^{19 20}</p> <p>Oral HPV infections... were as likely as cervical infections to persist to 6 months.²¹</p>	<p>4. HPV can usually be cleared by the immune system but occasionally, it stays in the cells of the affected area for several years. This persistent infection causes changes in the cells that may lead to cancer.</p>	
<p>Even if HPV causes cells to become abnormal, it usually takes up to 10 years for cervical cancer to develop.*</p>		<p>-----</p> <p>A few women were concerned about the length of time between HPV infection and cervical cancer developing.... Information that the transition from HPV infection to cervical cancer was slow was an</p>	<p>Progression from mild to severe dysplasia was at a constant low rate of 1% per year for up to 10 years... At 2 years, 29% of mild dysplasia was still present or had progressed. Progression from mild to moderate or worse was 11% (95% CI = 10%-12%) in 2 years. At 10 years, over half of the women with either mild or moderate dysplasia had not progressed to a more severe lesion.²²</p>	<p>5. Even if HPV causes cells to change, it can take an average of 10 to 15 years for cancer to develop.</p>	

<p>You get HPV from having sex or intimate sexual contact with another person.*</p>	<p>Large numbers of survey respondents in the public and professional groups (91.6% and 70.7% respectively) believed the information that HPV infection can be transmitted through any intimate genital contact was an important message.</p>	<p>important actor which reduced some women's anxiety.⁸</p> <p>-----</p> <p>Some women expressed surprise at the information that HPV could be passed by skin to skin genital contact.⁸</p>	<p>The natural history of oral HPV infection mimics that of genital HPV in general.²³ <u>HPV fact sheet:</u> HPV is spread from one person to another during sex and skin to skin contact.⁴</p> <p>The virus is... transmitted via contact with infected body areas. This includes vaginal, anal and oral sexual behaviours.²⁰</p> <p>Possible modes of transmission include oral-genital contact, oral-anal contact, and oral-oral contact.¹⁹</p> <p>HPV is very common and can be transmitted through vaginal and anal sex. However, oral sex, kissing, and genital-to-genital contact or genital-to-hand-to-oral contact can also allow for transmission to occur.⁶</p>	<p>Clinicians find it hard to explain about the sexually transmitted nature of HPV infection, and worry about causing patients distress and worry.</p> <p>"..Finding out it is sexually transmitted is going to be upsetting for them – with connotations of feeling guilty about previous relationships, or angry at their partners."</p> <p>"I don't want to go down the route of telling the patient it is to do with sexual transmission because it can open a huge can of worms. It's like HIV, herpes etc. – because they don't know enough about it."</p> <p>"The hardest thing to explain is that it is a sexually transmitted disease but it's not the same thing as having Chlamydia. Worries they don't understand."</p> <p>"Even if they only have a single partner throughout their life, they can still have HPV and then it would open a can of worms." There is debate whether condom use actually makes any difference; but you can't say to a sexually active women that no, don't do this.</p> <p>"Wearing a condom makes no difference, it doesn't give any more protection."</p>	<p>6. You get HPV from having sex or intimate contact with another person.</p>
<p>Although it is a good idea to use condoms to protect your sexual health, they do not provide complete protection against HPV.</p>	<p>In interviews with vaccination-aged girls and their parents, some parents who declined the vaccine believed that barrier methods of contraception offered as good a level of protection against HPV as the vaccine.</p>	<p>Among 27 estimates from 20 studies, there was no consistent evidence that condom use reduces the risk of becoming HPV DNA-positive.²⁴</p> <p>-----</p> <p>Some women asked questions about the ability of condoms to protect against contracting HPV... Others assumed the condoms would be effective (as they are against STIs in general).⁸</p> <p>Some thought condom use negated the use for a vaccine.³ HPV infections clear within 2 years in more than 90% of individuals.²⁵⁻²⁷</p>	<p><u>HPV fact sheet:</u> Condom use does not totally protect against HPV.⁴</p> <p>... if HPV is present on uncovered skin or non-genital mucosal sites such as the oropharynx, transmission is possible... Clearance of HPV and regression of HPV-associated penile lesions and CIN has been reported with consistent condom use. However, this recommendation may not be practical for individuals in long-term, monogamous relationships, or relevant to HPV infection in the oropharynx.⁶</p> <p>"...something I did 25, 30 years ago came back to haunt me... "I wanted to find out who the heck it was who gave it to me."²⁸</p> <p>...there is a significant "latent period" ... which makes it impossible to define when, where and from whom any individual acquired a high-risk HPV.²⁰</p>	<p>The main issue is that people might start blaming themselves.</p> <p>"The important message to get over to patients is that we are all exposed to viruses during normal sexual or social interaction and that their immune system has not knocked this one out...and a whole series of</p>	<p>7. Although it is a good idea to use condoms to protect your sexual health, they do not provide complete protection against HPV.</p>
<p>There is no way of knowing when you got HPV or who you got it from.</p>					<p>8. Most people never know they had HPV because there aren't any symptoms so there is no way of knowing when you got</p>

<p>Having a diagnosis of HPV-related cancer doesn't mean you or your partner have been unfaithful or promiscuous.</p>			<p>It is important to tell patients that a diagnosis of OPC does not suggest recent HPV infection, that exposure may have occurred many years before, and that a diagnosis does not suggest recent infidelity, either on the part of their partner or themselves.⁵</p> <p>It is impossible to know for certain from whom or when one initially acquired HPV because most people do not know they have it.⁶</p> <p>The patient may have been exposed in another relationship months, years, or decades earlier, and the infection may have been dormant ("hidden") in the meantime.⁶</p> <p>It is important to remind patients that being diagnosed with HPV-OSCC does not imply that either partner was/is unfaithful.... Or that either partner has a "risky" sexual past... Although HPV-OSCC is associated with sexual behaviours and a higher number of sexual partners... it only takes one partner who is infected to acquire the infection.¹⁹</p> <p>Because HPV is a common infection and usually acquired early in life it doesn't imply infidelity or promiscuity; there should be no shame or blame associated with it.^{6 19}</p>	<p>things have happened since which have led to you having a tumour."</p> <p>"It is very likely that their partners also have it, not necessarily through them."</p> <p>"..so it's the last thing he's going to do is tell is patients it's their previous sexual activities which caused this...it can open a huge can of worms..."</p> <p>Care must be taken not to imply that it's the patient's fault and that they were promiscuous when they may well not have been. Having the virus does not necessarily mean that you have had lots of sexual partners.</p> <p>"It is a marker of sexual activity not sexual promiscuity"</p> <p>"The younger ladies think it's associated with promiscuity whereas the older ladies tend not to ask anything."</p> <p>"Did I give it to my husband, or did he give it to me? One patient was distraught having been told that he had given it to his wife, or vice versa."</p>	<p>HPV or who you got it from.</p>
<p>You can get infected from one sexual experience, but you are more likely to get HPV the more sexual partners you have and the more partners they have had.*</p>	<p>Nearly all of the members of the public surveyed (98.8%), and half of the healthcare professionals (54.4%) believed the statement "Having many sexual partners increases the risk of HPV infection" to be false.</p>	<p>HPVs infect epithelial cells in genital mucosa (alphapapillomaviruses only), oral mucosa or skin (representatives of all five genera)²⁹</p> <p>-----</p> <p>A common belief was that girls or women only contracted HPV if they slept around.³</p> <p>A few acknowledged that a partner's sexual history was relevant... In studies of women who had been tested for HPV, the underlying concern about disclosure centred on embarrassment and shame about the promiscuity attached to an STI and the fear of being sexually rejected... Some participants expressed anger</p>	<p>Risk factors for HPV-positive OPSCC include specific sexual behaviours related to numbers of oral (>5) and vaginal (>25) sexual partners.³⁰</p> <p>As with all other sexual behaviours, safer sexual practices and fewer sexual partners may be considered the mainstay of prevention.⁶</p> <p>Female HPV prevalence and acquisition have been positively associated with women's estimates of their male partners' lifetime number of partners or not knowing a male partner's prior sexual history.³¹</p> <p>It is clear that major risk factors for acquiring genital HPV involve sexual</p>	<p>Clinicians explain that you only need one sexual relationship to be exposed to HPV, although risk factors are multiple partners or starting sex life at a young age.</p> <p>"In effect once is enough"</p> <p>I will explain that there has been an association with people who have multiple sexual partners or that homosexual practices have been associated with HPV.</p>	<p>9. Having a diagnosis of HPV-related cancer doesn't mean you or your partner have had sex with someone else.</p> <p><i>Deselected after consultation with expert group</i></p>

<p>HPV targets the moist red or pink tissue known as mucous membrane</p>	<p>and frustration when they learned that HPV could be contracted even if one had had only one sexual partner in one's lifetime.⁸</p> <p>HPVs infect epithelial cells in genital mucosa (alphapapillomaviruses only), oral mucosa or skin (representatives of all five genera).²⁹</p>	<p>behaviour, particularly multiple sex partners.³²</p> <p>HPVs can infect basal epithelial cells of the skin or inner lining of tissues and are categorized as cutaneous types or mucosal types... Mucosal types infect the lining of the mouth, throat, respiratory tract, or anogenital epithelium.^{13 33 34}</p>	<p>One clinical nurse specialist working with head and neck and gynae patients was drafting her own leaflet to answer patients' questions including,</p> <p>"...targets moist pink or red tissue known as mucous membrane."</p>	<p>10. HPV targets the moist red or pink tissue known as mucous membrane</p>
<p>Smoking may worsen your prognosis, especially if you continue to smoke after diagnosis.</p>	<p>Current smoking has associated with a significant increased risk of squamous cell carcinoma.³⁵</p>	<p>The New Zealand HPV project http://www.hpv.org.nz/patient-information/hpv-in-perspective Accessed 04/03/2016</p> <p>Tobacco use at the time of diagnosis, or tobacco use that continues during treatment in patients with HPV-HNC are independently associated with worse prognosis,^{36 37} particularly in response to adjuvant radio-therapy.³⁸</p> <p>The biological behaviour of HPV-positive tumours may be altered by tobacco use.^{38 39}</p>	<p>If you are a smoker it is more likely to persist and cause abnormalities and cutting down on smoking can help get rid of the virus.</p> <p>"We strongly emphasise the no smoking every time they come."</p>	<p>11. Smoking may worsen your prognosis, especially if you continue to smoke after diagnosis.</p>
<p>There is no need to inform previous partners about your HPV status, and it's up to you whether to tell your current partner.</p>		<p>HPV positive patients should be counselled on the benefits of cessation.⁶</p> <p>...there is no evidence to support partner notification of HPV-positive patients or referral for clinical evaluation of their partners. The current ethically acceptable approach to STI testing supports this notion, as notification of partners may cause unnecessary and unethical distress given the absence of clinical standards of treatment. However patients may benefit from having an informed discussion with their partner about their diagnosis, which can foster partner support.^{6 19 40}</p>	<p>Patients asked who they might have caught it off, do they need to tell them, are they at risk of spreading it?</p> <p>"It's not the same as having an STI like chlamydia or gonorrhea. You don't need to do the contact tracing. It's not the same in that way."</p>	<p>12. HPV is a sexually transmitted infection but, unlike other STIs, there is no need to inform previous partners about your HPV status. It is up to you whether you tell your current partner.</p>
<p>It's too late for you or your partner to be vaccinated but it's a good idea for your</p>	<p>Among women aged 15-25 years not previously infected with vaccine type HPV strains, prophylactic HPV vaccination appears to be highly efficacious in preventing HPV</p>	<p>"Long term partners do not appear to be at increased risk of HPV infection, and there is no evidence to support changes in intimate/sexual behaviour with a current partner."⁴¹</p> <p>Vaccination has no role for adults who have already been sexually active for a number of years, or someone who has already been infected with HPV and no role in cancer treatment. It is recommended for</p>	<p>Patients are interested to hear about the vaccine, even though it is not immediately relevant to them. They see it as a public health opportunity to help protect</p>	<p>13. There is no treatment for HPV infection but there's now a vaccination for</p>

children/ grandchildren	infection and precancerous cervical disease ⁴²	adolescents and young adults, ideally prior to the onset of sexual activity. ⁴³	daughters/granddaughters, and wonder about a vaccination for boys.	people aged 9-26. It's a good idea for young people to be vaccinated before they become sexually active.
HPV can be passed without penetrative sex but it does require intimate physical contact	The primary mode of human papillomavirus (HPV) transmission is through penetrative sex; however, there is evidence of other modes of transmission... Human papillomavirus DNA was detected in the genital tract of female virgins, with prevalence estimates ranging from 0% to 51.1%. HPV transmission from hands to genitals or genitals to hands was reported for both sexes and heterosexual couples. ⁴⁴	The available vaccines cover the HPV genotypes found most commonly in the oral mucosa but their protective effect against oral/oropharyngeal cancer, especially in individuals who have already been exposed to HPV, remains to be demonstrated. ⁶	"I would pay for my son to have the vaccination as soon as he is 12/13." "Patients mention about teenage girls having a vaccination – but what about boys? It's giving the wrong message and letting boys off the hook again when they should have responsibility too".	14. HPV can be passed without penetrative sex but it does require intimate physical contact
HPV can be passed by the hands from one intimate part of the body to another	HPV transmission from hands to genitals or genitals to hands was reported for both sexes and heterosexual couples. ⁴⁴	HPV is sexually transmitted, but penetrative sex is not required for transmission. Skin-to-skin genital contact is a well-recognized mode of transmission. ⁴⁵		15. HPV can be passed by the hands from one intimate part of the body to another
About two thirds of oropharyngeal cancers (which can involve the tonsils, tongue base and soft palate are caused by HPV	...whereas in Europe there was a steady increase in HPV prevalence across all time frames, reaching nearly 50% most recently, in North America HPV prevalence appears to have plateaued over the past decade at about 65%. ⁴⁶ Overall HPV prevalence in OPC increased significantly over time: from 40.5% (95% CI, 35.1-46.1) before 2000, to 64.3% (95% CI, 56.7-71.3) between 2000 and 2004, and 72.2% (95% CI, 52.9-85.7) between 2005 and 2009 (p < .001). Prevalence increased significantly in North America and Europe, and the significant gap between them that existed before 2000 (50.7% vs 35.3%, respectively, p = .008) has now	HPV is very common and can be transmitted through vaginal and anal sex. However, oral sex, kissing, and genital-to-genital contact or genital-to-hand-to-oral contact can also allow for transmission to occur. ⁶ Almost 70% of all tonsillar and oropharyngeal carcinoma cases are HPV-positive in economically developed countries. ⁴⁸ According to recent data, 3 of 4 newly diagnosed oropharyngeal carcinomas are HPV-positive (HPV+); if current trends continue, HPV+ OPSCC prevalence is expected to overtake that of cervical cancer by the year 2020. ⁴⁹	"Head & Neck has become more relevant now that it's been associated with HPV infection and also with better prognosis, so you're giving people potentially useful information."	16. About two thirds of oropharyngeal cancers (which can involve the tonsils, tongue base and soft palate are caused by HPV

Oral HPV is usually transmitted by oral sex.

There is some evidence that HPV can be transmitted by “deep” or “French” kissing.

disappeared (69.7% vs 73.1%, respectively, $p = .8$)⁴⁷
Cancer of the oropharynx was associated with having a history of six or more lifetime sexual partners [OR = 1.25, 95% confidence interval (CI) 1.01, 1.54] and four or more lifetime oral sex partners (OR = 2.25, 95% CI 1.42, 3.58). Cancer of the tonsil was associated with four or more lifetime oral sex partners (OR = 3.36, 95% CI 1.32, 8.53), and, among men, with ever having oral sex (OR = 1.59, 95% CI 1.09, 2.33) and with an earlier age at sexual debut (OR = 2.36, 95% CI 1.37, 5.05). Cancer of the base of the tongue was associated with ever having oral sex among women (OR = 4.32, 95% CI 1.06, 17.6), having two sexual partners in comparison with only one (OR = 2.02, 95% CI 1.19, 3.46) and, among men, with a history of same-sex sexual contact (OR = 8.89, 95% CI 2.14, 36.8).⁵⁰

‘The increase in HPV and oral cancers is presumed to be associated with changes in sexual behaviour including oral sex.’³⁴

‘Oral HPV infection and/or OPC are not markers of sexual promiscuity or abnormality. Oral sex is a normal sexual activity; in the US, 80% of sexually active people between 15-44 have had oral sex with an opposite sex partner (CDC fact sheet), whereas in the UK >70% aged 16-44 reported oral sex with an opposite sex partner in the last year.’⁵¹

‘Affected patients want answers to questions that clinicians may find difficult to answer, because these questions often fall far outside the remit of traditional consultations held in head and neck clinics.’⁵

Oral sex and open-mouthed kissing are associated with the development of oral HPV infection.⁵²
“Can you get it... You know, just from kissing someone?”²⁸

‘...baseline oral HR HPV status was significantly predicted by the spouse’s oral HR HPV status at baseline ... Here, we found that oral sex habits did not predict the outcome of oral HPV of either spouse ...these results suggest that the oral-to-oral route might be an important mode of HPV transmission.’²³

Some people with oral HPV infection report never having performed oral sex.^{5 52 53}
Autoinoculation from male genitals to oral cavity is a possible route of transmission.⁵⁴

It is also possible that HPV can be transmitted by deep kissing but data are currently inconclusive.⁵²⁻⁵⁶

Modes of transmission that have been proposed include kissing, other mouth-to-mouth contact, autoinfection, and perinatal contact.⁵⁷

We usually start by saying this has been caused in the past by smoking and drinking related factors, but now we’re finding over the past few years that this is actually caused by a sexually transmitted virus. It is important to publicise that oropharyngeal cancers can be caused by oral sex. This is a very sensitive topic.

[Patients ask,] “I have never smoked – why have I got this?”

“HPV is usually transmitted to the mouth by close sexual contact.”

“If the couple haven’t engaged in oral sex, then it could be from a past relationship – this can be awkward.”

“She had 2 patients – one said he felt vulgar, embarrassed about his lifestyle and it made him feel dirty and the other said the same.”

Clinicians are concerned that their patients worry about transmitting HPV to their wife/husband/children. They ask that if it is in their mouth, is it something they can transfer from kissing?

“Having one relationship, even a kiss, not even a sexual relationship, could result in transmission. How can you tell a whole generation of people that they can’t kiss, even though you might be able to tell them not to be promiscuous, you can’t tell them not to fondle or kiss.”

17. Oral HPV infection can be transmitted by oral sex as well as other forms of close sexual contact

18. There is some evidence that HPV may be transmitted by “deep” or “French” kissing but there is no evidence of transmission by sharing drinks or kissing on the cheek

<p>You have a better prognosis if you have HPV because this type of cancer responds better to treatment [Head and neck cancers only]</p>	<p>Patients with HPV-positive high grade non-squamous cell carcinoma had a lower risk of dying and a lower risk of recurrence than HPV-negative HNSCC patients. This review suggests that there is an improved overall and disease free survival for patients with HPV-positive tumours, but that this effect may be site specific to cancers of the oropharynx.⁵⁸</p> <p>Both progression-free survival and disease-free survival were significantly improved in HPV-positive HNSCCs. HPV-positive HNSCCs and OPSCCs patients have a significantly lower disease specific mortality and are less likely to experience progression or recurrence of their cancer than HPV-negative patients; findings which have connotations for treatment selection in these patients.⁵⁹</p>	<p>“Once I knew they were looking for [HPV] and that it would be beneficial for me, I hoped that I would have it”²⁸</p> <p>HPV positivity is inversely associated with molecular markers of poor prognosis,^{60 61} significantly reduced risk of recurrence,⁶² and improved disease-free and progression-free survival.^{39 59 63} HPV-HNC responds better to primary surgical therapy,⁶⁴ adjuvant radiotherapy⁶⁵ and/or chemo-radiotherapy protocols.^{66 67}</p> <p>Be positive – “We think your cancer is related to a viral infection which is good news”⁵</p> <p>HPV-related OPC has a much better prognosis than non-HPVOPC. It is associated with a better response to treatment resulting in significantly reduced risk of recurrence and improved disease-free and progression-free survival, as well as a significant reduction in disease-specific mortality.^{19 20 43}</p>	<p>“People with this tend to react better to treatment.”</p>	<p>19. You are likely to have a better outcome if your cancer has been caused by HPV because HPV-driven cancers respond better to treatment than other cancers. [Head and neck cancers only.]</p>
<p>If you have HPV, your partner is likely to have HPV too. There is no way to know who gave it to whom.*</p>	<p>This was a particularly important point for women who were HPV positive; they feared they would be thought to have been unfaithful, feared their partner had been once their HPV status was identified. (interviews with women in the UK cervical screening programme)</p>	<p>The majority ...felt uninformed about whether precautions should be taken to safeguard their partner from HPV.⁶⁸</p> <p>It is impossible to know for certain from whom or when one initially acquired HPV because most people do not know they have it.⁶</p>	<p>This causes distress among patients. Many wonder whether their partner has been unfaithful (especially if they have had been with the same partner for a long time).</p> <p>“Patient’s wife has cervical cancer and he is convinced she gave it to him. I explain that it could be either way.”</p>	<p><i>Deselected after consultation with expert group</i></p> <p><i>Content overlaps with message 8</i></p>
<p>Partners of patients with HPV-related cancers may be at slightly increased risk, but these are still rare</p>	<p>Some women understood that the transmission of HPV could have occurred prior to their present relationship – thus making it difficult to determine whether any infidelity had occurred.⁸</p>	<p>Partners who have been together tend to share HPV. This means the partner likely has HPV already. There is no way to know if the partner gave the patient HPV or vice versa. Because HPV is a common virus and is usually acquired early in life, having an HPV infection does not imply infidelity. It is important to remember that there should be no shame or blame associated with a diagnosis of HPV.⁶</p> <p>Partners of patients with HPV-OSCC may have slightly higher rates of HPV-associated cancers such as anal, penile cervical and/or oropharyngeal than the general population, but these are still rare cancers and the</p>	<p>“The wife was so stressed out because she thought she had caused her husband’s cancer...and another couple where the wife was distraught because she thought it was due to her husband being unfaithful to her.”</p> <p>“Even if they have an open and honest relationship with their partner, you still have that culture of blame of how has it happened, have they been sleeping with somebody else.”</p> <p>There are many worries from patients about passing on HPV.</p> <p>“Don’t worry about your kids, you’re not going to spread it”</p>	<p><i>Deselected after consultation with expert group</i></p>

			<p>cancer. It is recommended that these individuals use regular dental appointments as important tools for detecting any high-risk HPV-related changes in the oral and oropharyngeal mucosa.⁶</p> <p>Female partners should undergo routine cervical screening according to current guidelines.⁷⁰</p> <p>A clear dose-response relationship exists between the intensity and duration of tobacco exposure and individual risk of developing HNC.²⁰</p> <p>Behavioural modification, such as cessation of alcohol and tobacco use, should be encouraged because they are not only health-enhancing strategies but have been found to significantly decrease the odds of oral HPV persistence.^{6 21}</p>		
<p>If he/she is a smoker, stopping smoking will reduce your partner's risk of developing HPV-associated cancer.</p>					<p><i>Deselected after consultation with expert group</i></p>
<p>HPV vaccination also provides some protection against other cancers such as <i>penile, vulvar, vaginal, anal and head and neck</i>.</p>	<p>Women interviewed in the context of cervical screening questioned whether cancers other than cervical cancer can be caused by HPV, particularly whether men are more at risk from other HPV related cancers.</p>	<p>40% of vulvar, 60% of vaginal and 80% of anal carcinoma may be avoided by prophylactic vaccines against HPV 16/18¹⁶</p> <p>Oral infection with HPV is a significant independent risk factor for oral squamous cell carcinoma.¹¹</p> <p>-----</p> <p>Some participants questioned whether HPV can cause other cancers.⁸</p>		<p>The media publicity related to the vaccination programme has been useful to start people talking.</p> <p>"Tells them about the vaccination programme for young girls to reduce cervical cancer – and she would definitely vaccinate boys as well as HPV is implicated in so many cancers".</p>	<p><i>Deselected after consultation with expert group</i></p>
<p>This is the same cancer that Michael Douglas had</p>			<p>Suggested response to questions about MD: "Yes, Michael Douglas also had an HPV-related cancer and has done very well. Yet everyone's cancer is different and a personalised plan for the treatment of your specific cancer will be put together to achieve the highest cure rate with the least morbidity."⁴³</p> <p>In anal cancers, HPV-DNA is detected in 88–94%.¹³</p> <p>High-risk human papillomavirus (HPV) types can be detected in 80-90% of all anal SCC cases, making it second only to cervical cancer in the closeness of its association with this virus. HPV-16 can be detected in ~90% of HPV-positive cases of anal SCC.⁷²</p>	<p>Patients don't seem to be aware of HPV as such, but will ask if this is sexually transmitted like Michael Douglas.</p> <p>"Useful publicity to get people talking."</p>	<p><i>Deselected after consultation with expert group</i></p>
<p>Nearly all anal cancers are caused by HPV</p>		<p>This review provides evidence that anal HPV infection and dysplasia are common in women... The incidence of anal cancer continues to grow in all women, especially those living with HIV.⁷¹</p>		<p>" in anal cancer is it useful to give information around this as standard or are we just raising another issue which doesn't really help them?"</p>	<p>20. Nearly all anal cancers are caused by HPV</p>

Anal HPV can be transmitted by anal sex or by hand transmission or intimate skin-to-skin contact

In women it is easy for HPV to be transferred between the genital and the anal area

HPV has been detected in... 80-90% of anal cancers, with HR HPV types 16 or 18 detected in about 70% of cervical and 80% of anal cancers.⁷³

Current data indicate that HPV infection is potentially associated with 90%–93% of anal cancers⁷⁴

HPV is sexually transmitted, but penetrative sex is not required for transmission. Skin-to-skin genital contact is a well-recognized mode of transmission.⁴⁵

Among men who were not exclusively heterosexual and women, receptive anal intercourse was related strongly to the risk of anal cancer (OR, 6.8 [95% CI, 1.4 –33.8] and OR, 2.2 [95% CI, 1.4 –3.3], respectively).⁷⁵

Sixteen events involved transmission between different genital sites, 2 involved anal-to-genital transmission, and 3 involved genital-to-hand transmission.... Among the 5 couples with penis to-anus transmission, 4 reported anal intercourse during the corresponding time period.... Sexual transmission also involved the scrotum, the anus of women, and the hands of both sexes.... Transmission through nonpenetrative sexual contact was demonstrated between the female anus and the scrotum, as well as the female hand and male genitals. ⁷⁶

HPV is sexually transmitted, but penetrative sex is not required for transmission. Skin-to-skin genital contact is a well-recognized mode of transmission.⁴⁵

Some clinicians find this topic difficult to discuss with patients. Patients worried about stigma related to anal sex and what the clinician might think of them. It can be problematical with partners being present at a consultation.

“Obviously there is the topic of homosexuality and I find it quite difficult to ask if they have had anal intercourse, but having said that I don’t routinely ask the women because I find that difficult as well.”

“Whereas more historically with cervix cancer it is associated with a virus, can be associated with sexual contact – it’s within the vagina, that’s normal sexual practice and therefore I’m normal and it’s nothing that will have stigma attached to it, whereas those with an anal cancer may feel there is an element of stigma.”

“He has had one patient with tumours and who has had repeated tumours and been tested for HPV (positive) and is adamant that this is not related to anal intercourse and is quite bitter and angry...so they just leave it.”

Men and women need different information.

“He tells the women they have cancer and about treatment and follow up, which includes vaginal examinations so explains that the two things can be linked and they are at slightly increased risk.”

“I think you could possibly do with separate literature for men and women. We need to be careful as the one thing that does upset people, especially the women, is if they are

21. Anal HPV can be transmitted by anal sex or by hand transmission or intimate skin-to-skin contact

22. In women it is easy for HPV to be transferred between the genital and the anal area

<p>You can't get cervical cancer without having HPV first</p>	<p>The strength of the association between HPV persistence and cervical neoplasia increased with increasing grade of cervical disease. This trend supports the view that CIN 1 represents active, mainly transient HPV infection and has high rates of occurrence and regression among sexually active women. In contrast, long-term HPV positivity is clearly associated with neoplastic transformation.¹⁸</p>	<p>About 1000 women with histologically verified invasive cervical cancer were recruited from 22 countries around the world.... HPV-DNA was detected in 99.7% of the tumors, leading to the conclusion that HPV is a necessary cause of cervical cancer.¹³</p>	<p>lumped in a group with HIV positive homosexual men or drug addicts.” “You can't get cervical cancer without having it first.” “Cervical cancer and abnormalities of the cervix are generally related to the HPV virus.”</p>	<p>23. You can't get cervical cancer without having HPV first</p>
<p>Nearly half of vulvar and two thirds of vaginal cancers are caused by HPV</p>	<p>‘...HPV appears to be present in 65%–75% of vulvar and vaginal squamous cell cancers.’⁷⁷</p>	<p>Between 64 and 91% of vaginal cancers and 82 and 100% of VAIN-3 lesions are HPV-DNA positive.¹³</p> <p>Cancers of the vulva and of the penis have been associated to HPV. The tumors diagnosed in young individuals are usually of histological types called basaloid or warty; the majority (60–90%) are positive for HPV and their pre-neoplastic lesions are also strongly associated with HPV.¹³</p> <p>Current data indicate that HPV infection is potentially associated with... 40%–64% of vaginal cancers, and 40%–51% of vulvar cancers.⁷⁴</p>		<p>24. Nearly half of vulvar and two thirds of vaginal cancers are caused by HPV</p>

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